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Article

Characteristics, Treatment, Outcomes, and Cause of Death of Invasively Ventilated Patients with COVID-19 ARDS in Asir Central Hospital: A Retrospective Single Center Experience

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Abstract: COVID-19's emergence caused devastating consequences worldwide, with high morbidity and mortality rates. Critically ill patients face challenges like acute respiratory failure requiring intensive measures, including mechanical ventilation. Mortality rates in ARDS are influenced by various factors. This paper focuses on describing characteristics, management, and survival factors of mechanically ventilated ICU patients in Saudi Arabia. A retrospective, noninterventional approach, we reviewed charts and data of all COVID-19-infected patients who were admitted to the ICU, particularly those requiring ventilation support at Aseer Central Hospital (ACH) in the southern region of Saudi Arabia. The study enrolled 594 COVID patients, mean age 60.5 ± 17.3 years, with 67% males. Approximately half had O negative blood group. Common chronic health issues: diabetes mellitus (35.5%). Positive culture findings in almost 340 (57.2%), with Klebsiella pneumoniae being the most isolated microorganism (45.5%). Significant correlations found with mortality: age, blood culture, Rh positive group (P<0.05). This study provides critical insights into the clinical and epidemiological aspects of COVID-19 patients admitted to the ICU in a Saudi Arabian hospital. The research underscores the impact of age, comorbidities, laboratory markers, and treatment interventions on patient outcomes, emphasizing the significance of tailored management strategies in this context.

Keywords: COVID-19; ICU; Mechanical Ventilator; ARDS

1. Introduction

The advent of coronavirus disease 2019 (COVID-19) in late 2019 marked a pivotal moment with far-reaching and devastating consequences, giving rise to an unprecedented surge in morbidity and mortality rates globally. This highly infectious and contagious virus, classified within the Coronaviridae family, swiftly traversed geographical boundaries, casting its shadow on every corner of the world and precipitating profound challenges for economies and healthcare systems alike (1,2). In March 2020, the Kingdom of Saudi Arabia bore witness to its initial case of COVID-19, a significant milestone that prompted swift and decisive actions by the Saudi Ministry of Health. Faced with the emergent threat, the ministry escalated preparations and implemented robust precautions to effectively control the outbreak and cater to the medical needs of the affected population. The strategic response included the allocation of resources across a network of more than 25 hospitals, boasting a collective capacity of 8,800 beds. This infrastructure encompassed not only 8,000 intensive care unit (ICU) beds but also 2,200 isolation beds dedicated to suspected and quarantined cases, illustrating a comprehensive and systematic approach to patient care during the pandemic (3,4,5).

The magnitude of this concerted effort was paramount in successfully managing the expected influx of infected individuals. By proactively expanding healthcare resources and adhering to meticulous treatment guidelines, Saudi Arabia positioned itself to navigate the complexities of the pandemic and provide optimal care to those affected by COVID-19. This multifaceted response not only exemplifies the nation's commitment to public health but also serves as a noteworthy model for countries grappling with the challenges posed by the relentless spread of the virus.

The manifestation of coronavirus disease 2019 (COVID-19) is characterized by a spectrum of flulike symptoms, encompassing fever, cough, myalgia, and gastrointestinal issues, as documented in various studies (6). While a substantial proportion of cases present with mild symptoms, it is noteworthy that approximately one-quarter of affected individuals undergo a more severe clinical trajectory, potentially progressing to the challenging stage of acute respiratory distress syndrome (ARDS). Alarmingly, about 5% of cases escalate to a critical condition, requiring specialized and intensive medical intervention (7). Critically ill COVID-19 patients confront a constellation of unique challenges that significantly compound the complexity of their medical condition. These challenges include but are not limited to acute respiratory failure characterized by resistance hypoxemia, coagulopathy, multi-system failure, and discernible alterations in both immunity and inflammatory responses (8). The intricate nature of these complications underscores the importance of a meticulous and attentive approach to patient management. As healthcare providers navigate the multifaceted landscape of critical COVID-19 cases, they are compelled to address the intricate interplay of diverse physiological systems, necessitating a comprehensive and nuanced care strategy to optimize patient outcomes. The heightened complexity of the clinical presentation underscores the critical need for a tailored and multidisciplinary approach to meet the distinctive challenges posed by severe manifestations of COVID-19.

The critical care management of patients grappling with severe manifestations of COVID-19 in the intensive care unit (ICU) necessitates the application of a multifaceted approach, incorporating not only generic measures but also drawing upon evidence-based medicine guidelines. One pivotal facet of this management involves the implementation of invasive mechanical ventilation (MV) as an integral therapeutic intervention, a strategy underscored by the evolving understanding of the disease dynamics and clinical outcomes (9). In the context of acute respiratory distress syndrome (ARDS), a condition encompassing diverse etiologies, mortality rates are influenced by an intricate interplay of various factors. Patient-specific variables, including co-morbidities, advanced age, the severity of hypoxemia, and the presence of co-existing organ failures, collectively contribute to shaping the trajectory and ultimate outcomes in ARDS cases (10). The recognition of these nuanced determinants underscores the imperative for a personalized and comprehensive approach to critical care management. As clinicians navigate the complexities inherent in severe COVID-19 cases, the integration of evidence-based guidelines and a nuanced understanding of individual patient profiles becomes paramount. The incorporation of invasive mechanical ventilation aligns with the evolving landscape of critical care interventions, emphasizing a patient-centered paradigm to enhance the prospects of favorable clinical outcomes.

Despite the growing body of research on the clinical characteristics of intensive care unit (ICU) patients in Saudi Arabia, a notable research gap persists, particularly in the realm of those necessitating mechanical ventilation. This study endeavors to address this lacuna by delving specifically into the attributes and treatment modalities deployed for critically ill COVID-19 patients admitted to ICUs, with a meticulous focus on individuals requiring mechanical ventilation. The overarching goal is to contribute nuanced insights that not only augment the existing literature on severe COVID-19 cases but also shed light on the distinctive challenges and intricacies associated with the management of this specific subgroup.

2. Materials and Methods

Our comprehensive investigation delved deeply into the landscape of COVID-19 ICU admissions at Aseer Central Hospital (ACH), strategically situated in the southern region of Saudi Arabia. Adopting a rigorous retrospective, noninterventional methodology, we meticulously

scrutinized the medical charts and data of all patients who found themselves in the challenging realm of ICU admission due to COVID-19, with a particular emphasis on those necessitating crucial ventilation support. The ambit of our study extended to encompass hospitalized individuals aged 18 years and above, unequivocally diagnosed with COVID-19 through positive real-time polymerase chain reaction tests for SARS-CoV-2. Our data collection phase, spanning from August 2020 to April 2021, aimed to encapsulate a comprehensive snapshot of the evolving landscape of COVID-19 in the ICU setting. Following an assiduous process of data cleaning and the exclusion of cases not meeting our stringent inclusion criteria, our final study cohort manifested as a robust sample size, comprising 594 patients who intricately fit the specified criteria.

The granularity of our study extended beyond mere demographic details, as we meticulously gathered and organized data encompassing various facets of patients' profiles. This included indepth information on age, gender, nationality, and the intricate tapestry of comorbidities that can significantly shape the clinical trajectory of COVID-19 cases. A specific lens was directed towards unraveling the ICU profile, probing into the diverse interventions undertaken within this critical care milieu. The overarching goal was to unearth insights into patient mortality, a pivotal outcome of interest, and to explore potential associations with blood group types. Such a nuanced exploration demanded meticulous attention to detail and a robust ethical framework, as underscored by the approval granted by the Institutional Review Board at Aseer Central Hospital and our dedicated research committee. This ethical imprimatur ensures the integrity and compliance of our study with the highest standards of research governance, thereby bolstering the credibility and reliability of our findings.

2.1. Data analysis

Following the extraction of data, a meticulous and iterative process unfolded, involving thorough revision and coding procedures. Subsequently, the data seamlessly transitioned into the statistical software IBM SPSS version 26 (SPSS, Inc. Chicago, IL), marking a pivotal phase in our analytical journey. In adherence to robust statistical principles, two-tailed tests were employed for the spectrum of statistical analyses, with a predefined threshold of a p-value less than 0.05 deemed as statistically significant. The utilization of descriptive statistics emerged as a linchpin in our analytical framework, particularly for scale and continuous variables such as laboratory findings. Mean values accompanied by standard deviations provided a comprehensive overview of the central tendencies within this domain. In parallel, frequency distributions and percentages assumed prominence in delineating the landscape of categorical variables, encompassing personal data and blood group information.

The exploration of factors intricately linked to mortality among COVID-19 patients hinged on a multifaceted approach. Cross-tabulation, Chi-square tests, and exact tests, tailored to accommodate small sample sizes, collectively formed the methodological arsenal deployed for this nuanced investigation. Logistic regression analysis, a robust statistical tool, was judiciously applied to quantify the magnitude of the relationship between blood group, Rh factor, and the rate of patients' mortality. In the pursuit of a comprehensive understanding, the findings were not merely confined to statistical metrics but were thoughtfully presented graphically. Mortality rates, co-morbidities, medication usage patterns, culture findings, and other pertinent variables were graphically depicted, providing a visual narrative that enhances interpretability and fosters a nuanced appreciation of the multifaceted interplay within our dataset. This meticulous analytical journey, characterized by methodological rigor and a commitment to comprehensive reporting, underscores the reliability and depth of our research endeavor

3. Results

This study included 594 COVID patients admitted to Aseer Central Hospital (ACH) ICU. The patients' ages ranged from 18 to 103 years, with a mean age of 60.5 ± 17.3 years. Among them, 398 (67%) were males, and only 5 (0.8%) were smokers (Table 1).

Personal data	No	%
Age in years		
< 40	72	12.1%
40-59	193	32.5%
60-69	127	21.4%
70+	202	34.0%
Mean ± SD	60.5	± 17.3
Gender		
Male	398	67.0%
Female	196	33.0%
Smoking		
No	589	99.2%
Yes	5	.8%

Figure 1 displays the prevalence of chronic diseases among COVID patients admitted to the ICU at Aseer Central Hospital (ACH), Saudi Arabia. The most commonly reported chronic health issues in this study included diabetes mellitus (35.5%), dyslipidemia (13%), gastrointestinal diseases (5.1%), chronic kidney disease (5.1%), bronchial asthma (3.9%), and hypothyroidism (3.5%).

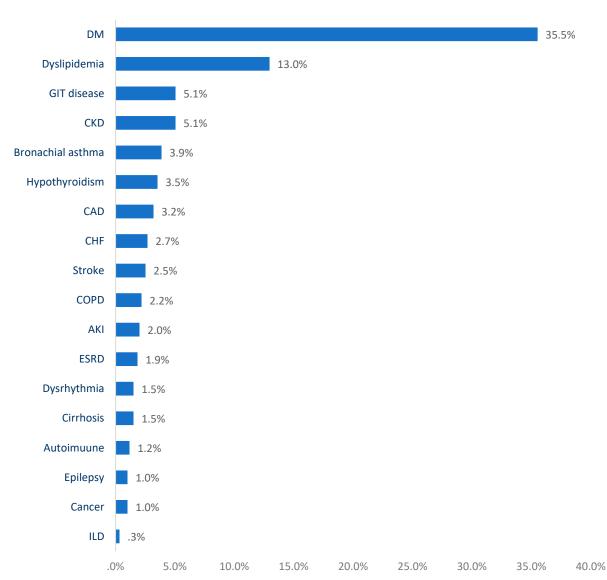


Figure 2 illustrates the medications administered to COVID patients admitted to the ICU. Upon admission, 95.3% received antibiotics, 87.4% were prescribed anticoagulants, 78.3% received steroids, 75.3% were given antiviral drugs, and 55.4% received analgesics. Monoclonal antibodies (MABS) were administered to 20.4% of patients, and 14.5% received antiplatelet medication. The average laboratory findings of COVID patients admitted to Aseer Central Hospital (ACH) ICU are displayed. The blood glucose level (BGL) was recorded as 276.7 ± 12.1 mg/dl, creatinine as 2.6 ± 1.9 g/dl, erythrocyte sedimentation rate (ESR) as 62.8 ± 35.6 , and C-reactive protein (CRP) as 60.4 ± 62.9 (Table 2). Regarding chest X-ray findings, tracheostomy tubes were observed in 235 (98.7%) patients, while only 6 (2.5%) displayed signs of consolidation (Table 2).

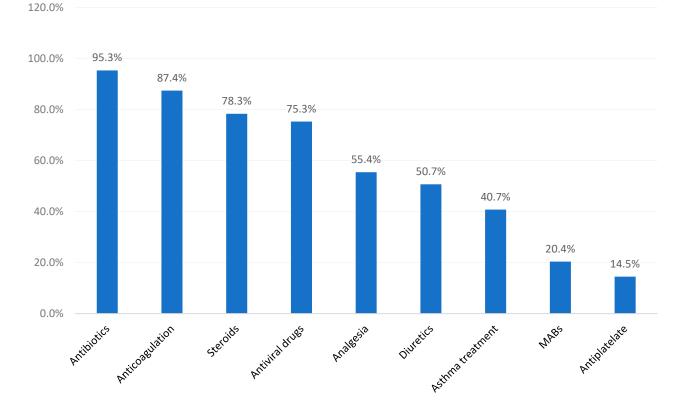


Figure 2. Medications taken by covid patients admitted to ICU, ACH, Saudi Arabia.

Table 2. Laboratory findings among covid patients admitted to ICU, ACH, Saudi Arabia.

Laboratory findings	Mean	SD
Creatinine	2.6	1.9
Hb	13.3	2.9
WBCs	19.9	12.0
Platelets	131.8	132.1
LDL	78.3	59.4
HDL	35.1	17.4
ESR	62.8	35.6
CRP	60.4	62.9
BGL	276.7	129.1
Total bilirubin	1.8	2.1

Almost 340 (57.2%) had positive culture findings. The most commonly isolated microorganisms included Klebsiella pneumoniae (45.5%), Acinetobacter (20.2%), Staphylococcus epidermidis (13.8%), Pseudomonas aeruginosa (6.2%), MSSA (5.6%), Staphylococcus haemolyticus (4.7%), and

Enterococcus faecium (4.7%) (Figure 3). Among the patients, 320 (54.6%) died, while 266 (45.4%) survived (Figure 4).



Figure 3. Culture findings among covid patients admitted to ICU, ACH, Saudi Arabia.

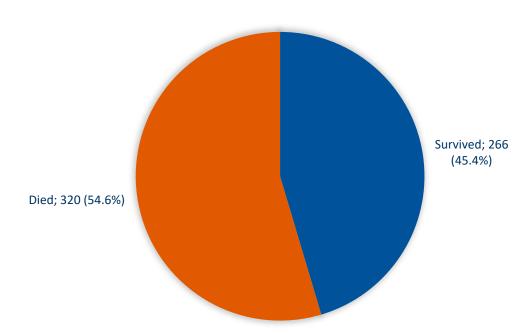


Figure 4. Death rate among covid patients admitted to ICU, ACH, Saudi Arabia.

In this study, a significant correlation was found between age and mortality, with 75% of patients aged 70 years or more succumbing to the disease compared to 22.5% of those under 40 years (P=.001). Patients with positive culture results also had a higher mortality rate of 67.4% compared to 37.7% for those with negative culture findings (P=.001), Other factors examined did not show significant relations with COVID patients' mortality (Table 3).

Table 3. Factors associated with mortality among covid patients admitted to ICU, ACH, Saudi Arabia.

Factors	Death				
	No		Yes		– p-value
	No	%	No	%	
Age in years					
< 40	55	77.5%	16	22.5%	
40-59	110	58.2%	79	41.8%	.001*
60-69	51	40.5%	75	59.5%	
70+	50	25.0%	150	75.0%	
Gender					
Male	171	43.6%	221	56.4%	.221
Female	95	49.0%	99	51.0%	
Smoking					
No	263	45.3%	318	54.7%	.510\$
Yes	3	60.0%	2	40.0%	
DM					
No	167	44.5%	208	55.5%	.578
Yes	99	46.9%	112	53.1%	
Culture					
No	157	62.3%	95	37.7%	.001*
Yes	109	32.6%	225	67.4%	

P: Pearson X^2 test; \$: Exact probability test; * P < 0.05 (significant).

4. Discussion

The imperative for intensive care unit (ICU) admission arises for a substantial proportion, approximately 25%, of hospitalized individuals grappling with the complexities of COVID-19. Recognizing and delineating the characteristics of high-risk individuals within this cohort assume paramount significance, as this discernment holds the key to mitigating mortality rates and elevating overall clinical outcomes in cases where the severity of the disease necessitates such intensive care interventions (11,12,13). Against this backdrop, the primary objective of this paper is to delve into the multifaceted realm of critically ill COVID-19 patients admitted to the ICU. More specifically, our focus extends to those within this critical subset who necessitate mechanical ventilation, a pivotal aspect of the intensive care landscape. Beyond a mere exploration of their characteristics and management nuances, our study embarks on a meticulous investigation into the factors intricately associated with patient survival within this highly specific and clinically intricate subgroup. By shedding light on the unique challenges and dynamics of this subset, we aspire not only to contribute to the enriched understanding of severe COVID-19 cases but also to offer nuanced insights that can inform tailored clinical strategies and ultimately enhance the outcomes for patients navigating the complexities of mechanical ventilation in the ICU setting.

In consonance with the trends delineated in various published studies, a substantial two-thirds majority within our study cohort comprises male individuals, a consistent observation mirrored in both national and international reports (14,15,16). An intriguing demographic nuance emerges as we delve into the age distribution of our intensive care unit (ICU)-admitted COVID-19 patients, where a noteworthy over 50% fall into the age bracket of 60 years or older. This demographic alignment resonates harmoniously with insights gleaned from an amalgamation of local and international studies, underscoring the predilection of older individuals toward more severe disease manifestations and complications. The predisposition of older patients to experience heightened disease severity and complications stems from a complex interplay of factors, including the presence of multiple comorbidities, compromised immunity, and an elevated susceptibility to various complications. This intricate web of vulnerabilities necessitates a nuanced understanding to facilitate targeted care strategies, ultimately culminating in improved outcomes for this particularly vulnerable population (14,17,18,19). Recognizing and comprehensively addressing these multifaceted risk factors not only enriches our understanding of the demographic dynamics within our study but also reinforces the imperative of tailored interventions to enhance the care trajectory and clinical outcomes for older COVID-19 patients admitted to the ICU.

Certainly, the intricate interplay between comorbidities and the trajectory of COVID infections is an undeniable facet of the evolving narrative surrounding this global health crisis (11,20). The surge in the prevalence of diabetes mellitus (DM) emerges as a formidable public health challenge, wielding a profound impact on long-term morbidity and mortality rates. Within the Gulf region, Saudi Arabia takes center stage, marked by a record of the highest diabetes-related deaths, as underscored by a pivotal study (21). The present investigation casts a revealing light on this landscape, uncovering that more than one-third of the patients under scrutiny bore the burden of a DM diagnosis. This prevalence aligns seamlessly with broader epidemiological evidence indicating that nearly half of individuals aged over 55 in Saudi Arabia grapple with type 2 diabetes mellitus (DM2). Importantly, this pattern resonates within the demographic contours of our patient age group (22). Significantly, our study's documentation of DM prevalence surpassed that reported in a study conducted in Alahsa (14), where approximately 20% of COVID patients exhibited DM, albeit with a focus outside the intensive care unit (ICU) context. Furthermore, our findings align with another influential cohort study on COVID patients, shedding light on the pervasive nature of DM in this cohort, with almost half of the patients grappling with this metabolic condition (17). This nuanced exploration not only enriches our understanding of the intricate relationship between DM and COVID-19 but also situates our findings within the broader landscape of regional and global studies, thereby contributing to a more comprehensive comprehension of the multifaceted challenges posed by the intersection of diabetes and severe COVID-19 manifestations.

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Numerous studies have delved into the intricate correlations between various laboratory parameters and the nuanced landscape of COVID-19. This expansive spectrum of parameters spans procalcitonin (PCT), lactic acid, the percentage of lymphocytes, D-dimer levels, C-reactive protein (CRP), erythrocyte sedimentation rate (ESR), and serum lactate dehydrogenase (LDH), each serving as a sentinel marker in the intricate interplay of the disease's pathophysiology (23,24). In the comprehensive laboratory assessments conducted on our patient cohort, conspicuous elevations were observed in the renal profile, hepatic enzymes, and inflammatory markers, with particular emphasis on heightened ESR and CRP levels. These findings not only echo the established literature but also fortify the strong association documented in prior studies between these inflammatory markers and COVID-19 mortality.

The elevated prominence of inflammation within the pathophysiological milieu of critically ill COVID-19 patients is further accentuated by our results, hinting at a substantial role played by these inflammatory markers in shaping the clinical trajectory of severe cases. Intriguingly, our findings also posit CRP as a potential independent player in influencing ICU admission, adding a nuanced layer to our understanding of the prognostic significance of this marker (25, 26, 19, 14). As standard practice aligns with the administration of dexamethasone to moderate to severely hypoxic COVID-19 patients, a strategy linked to improved outcomes, our study dovetails into this broader treatment paradigm. Furthermore, the amalgamation of antibiotics with hydroxychloroquine, as highlighted in a multicenter study, demonstrated reduced mortality in COVID-19 cases. Notably, adherence to health protocols and local guidelines was underscored by the remarkable statistic that over two-thirds of our studied patients received antibiotics and steroids, further contributing to the contextual richness of our findings (27,28,29).

This investigation has brought to the forefront a stark reality — a substantial mortality rate among severely ill COVID-19 patients admitted to the intensive care unit (ICU) and reliant on mechanical ventilation. Within this critical cohort, a discernible correlation between age and mortality has emerged, unveiling the nuanced impact of age on the trajectory of severe cases. Intriguingly, patients with positive culture outcomes, indicative of concomitant bacterial infections, displayed a notably elevated mortality rate, further accentuating the intricate web of factors influencing outcomes in critically ill patients. Notably, our findings harmonize with a spectrum of local studies conducted in Saudi Arabia and find resonance in a sweeping analysis encapsulating 178,568 COVID-19 deaths across 16 countries. This expansive investigation illuminates a striking reality - an approximately 8-fold higher mortality rate among individuals aged 55 to 64 and an astonishing over 62-fold higher incidence rate among those aged 65 or older, compared to their counterparts aged 54 or younger (16,18,19,24,25,30). This age-dependent vulnerability can be attributed to immunosenescence, a phenomenon where the performance of innate immune cells in older individuals undergoes compromise. This leads to a diminished efficacy in viral clearance and sets the stage for an imbalanced immune response (31). The profound implications of age on the mortality dynamics uncovered in our study find resonance in the broader global context, emphasizing the need for targeted interventions and heightened vigilance in managing severe cases, particularly in the elderly population grappling with the intricate challenges of immunosenescence.

5. Conclusions

In conclusion, this study provides critical insights into the clinical and epidemiological aspects of COVID-19 patients admitted to the ICU in a Saudi Arabian hospital. The research underscores the impact of age, comorbidities, laboratory markers, and treatment interventions on patient outcomes, emphasizing the significance of tailored management strategies in this context.

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supervision, Ali and Mushary; project administration, Ali and Mushary;. All authors have read and agreed to the published version of the manuscript.

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Informed Consent Statement: All subjects gave their informed consent for inclusion before they participated in the study. The study was conducted in accordance with the Declaration of Helsinki, and the protocol was approved by the Ethics Committee of Board at Aseer Central Hospital, IRB Log No.:REC-11-03-2022.

Conflicts of Interest: The authors declare no conflict of interest.

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